S-225N - 1 -

SEQUENCE LISTING

<110> LIN, LEU-FEN H

COLLINS, FRANKLIN D

DOHERTY, DANIEL H

LILE, JACK

BEKTESH, SUSAN

- <120> Glial Cell Line-Derived Neurotrophic Factor
- <130> S-225E Rev 070302
- <140> 08/182,183
- <141> 1994-05-23
- <150> 07/764,685
- <151> 1991-09-20
- <150> 07/774,109
- <151> 1991-10-08
- <150> 07/788,423
- <151> 1991-11-06
- <150> 07/855,413
- <151> 1992-03-19
- <150> PCT/US92/07888
- <151> 1992-09-17
- <160> 29

S-225N - 2 -

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Gln Ala Ala Ala Ser Pro Asp Asn
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<223> Xaa in position 2 is either Lys or Gln

S-225N - 3 -

-35

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gcc gcc gga cgg gac tct aag atg aag tta tgg gat gtc gtg gct gtc Ala Ala Gly Arg Asp Ser Lys Met Lys Leu Trp Asp Val Val Ala Val
                                                                                       99
                    -80
tgc ctg gtg ttg ctg cac acc gcg tct gcc ttc ccg ctg ccc gcc ggt Cys Leu Val Leu Leu His Thr Ala Ser Ala Phe Pro Leu Pro Ala Gly
                                                                                     147
               -65
                                       -60
                                                                                     195
aag agg ctt ctc gaa gcg ccc gcc gaa gac cac tcc ctc ggc cac cgc
Lys Arg Leu Leu Glu Ala Pro Ala Glu Asp His Ser Leu Gly His Arg
                                                                                     243
cgc gtg ccc ttc gcg ctg acc agt gac tcc aat atg ccc gaa gat tat
Arg Val Pro Phe Ala Leu Thr Ser Asp Ser Asn Met Pro Glu Asp Tyr
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-30

-25

														atc Ile		291
														cga Arg		339
														aga Arg		387
														tta Leu		435
														acc Thr		483
														gcc Ala 75		531
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														ttc Phe		627
														cta Leu		675
		tcc Ser								tgad	cctg	ggc t	ccag	gagad	et	725
gct	gtgta	att g	gcatt	ccto	gc ta	acact	gcga	a aga	aagg	ggac	caaç	gtto	ccc a	aggaa	aatatt	785
tgc	ccaga	aaa g	ggaag	gataa	ag ga	ccaa	agaag	g gca	gagg	gcag	aggo	ggaa	aga a	agaag	gaagaa	845
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<213> Rattus rattus

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S-225N - 5 -

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Ala Glu Asp His Ser Leu Gly His Arg Arg Val Pro Phe Ala Leu Thr -45 -35 -30

Ser Asp Ser Asn Met Pro Glu Asp Tyr Pro Asp Gln Phe Asp Asp Val -25 -20 -15

Met Asp Phe Ile Gln Ala Thr Ile Lys Arg Leu Lys Arg Ser Pro Asp -10 -5 -1 1

Lys Gln Ala Ala Leu Pro Arg Glu Arg Asn Arg Gln Ala Ala 5 10 15

Ala Ala Ser Pro Glu Asn Ser Arg Gly Lys Gly Arg Arg Gly Gln Arg 20 25 30 35

Gly Lys Asn Arg Gly Cys Val Leu Thr Ala Ile His Leu Asn Val Thr 40 45 50

Asp Leu Gly Leu Gly Tyr Glu Thr Lys Glu Glu Leu Ile Phe Arg Tyr 55 60 65

Cys Ser Gly Ser Cys Glu Ala Ala Glu Thr Met Tyr Asp Lys Ile Leu 70 75 80

Lys Asn Leu Ser Arg Ser Arg Leu Thr Ser Asp Lys Val Gly Gln 85 90 95

Ala Cys Cys Arg Pro Val Ala Phe Asp Asp Leu Ser Phe Leu Asp 100 105 110 115

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Gly Cys Ile

<210> 5

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<213> Homo sapiens

S-225N - 6 -

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									25					20		
		Asp				ttt Phe	Ile					Lys				101
		-15					-10					- 5				
Arg	Ser				Gln	atg Met				Pro					Asn	149
-1	1				5					10					15	
cgg Arg	cag Gln	gct Ala	gca Ala	Ala	gcc Ala	aac Asn	cca Pro	gag Glu	Asn	tcc Ser	aga Arg	gga Gly	aaa Lys	Gly	cgg Arg	197
				20					25					30		
			Arg			aac Asn		Gly					Ala			245
			35					40					45			
		Val				ggt Gly	Leu					Lys				293
		50					55					60				
	Phe					ggc Gly					Ala					341
	65		4.4			70					75					
Asp					Asn	tta Leu				Arg					Asp	389
80					85					90					95	40.5
Lys	Val	Gly	Gln	Ala	Cys	tgc Cys	aga Arg	Pro	Ile	gcc Ala	Phe	gat Asp	gat Asp	Asp	ctg Leu	437
+~~			~	100			~+-	.	105	a++				110		405
Ser	Phe	Leu	Asp	Asp	Asn	ctg Leu	Val	Tyr	His	Ile	Leu	aya Arg	Lys	His	Ser	485

S-225N - 7 -

gct aa Ala Ly		Cys				tga	ctcc	ggc	tcca	gaga	ct g	ctgt	gtat	t	536
gcattcctgc tacagtgcaa agaaag														562	
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Phe Ile		Ala	Thr	Ile	Lys -5	Arg	Leu	Lys	Arg -1	Ser 1	Pro	Asp	Lys	Gln 5	
Met Ala	a Val	Leu	Pro 10	Arg	Arg	Glu	Arg	Asn 15	Arg	Gln	Ala	Ala	Ala 20	Ala	
Asn Pro	o Glu	Asn 25	Ser	Arg	Gly	Lys	Gly 30	Arg	Arg	Gly	Gln	Arg 35	Gly	Lys	
Asn Arg	Gly 40	Cys	Val	Leu	Thr	Ala 45	Ile	His	Leu	Asn	Val 50	Thr	Asp	Leu	
Gly Let 55	ı Gly	Tyr	Glu	Thr	Lys 60	Glu	Glu	Leu	Ile	Phe 65	Arg	Tyr	Cys	Ser	
Gly Ser 70	c Cys	Asp	Ala	Ala 75	Glu	Thr	Thr	Tyr	Asp 80	Lys	Ile	Leu	Lys	Asn 85	
Leu Se	r Arg	Asn	Arg 90	Arg	Leu	Val	Thr	Asp 95	Lys	Val	Gly	Gln	Ala 100	Cys	
Cys Arg	g Pro	Ile 105	Ala	Phe	Asp	Asp	Asp 110	Leu	Ser	Phe	Leu	Asp 115	Asp	Asn	
Leu Val	l Tyr 120	His	Ile	Leu	Arg	Lys 125	His	Ser	Ala	Lys	Arg 130	Cys	Gly	Cys	
Ile															

536

S-225N - 8 -

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gaagttatgg gatgtcgtgg ctgtctgcct ggtgctgctc cacaccgcgt ccgccttccc
                                                                    120
gctgcccgcc ggtaagaggc ctcccgaggc gcccgccgaa gaccgctccc tcggccgccg
                                                                    180
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S-225N - 9 -

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223
ccgcgcgccc ttcgcgctga gcagtgactg taagaaccgt tcc
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<211> 12
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<223> Oligonucleotide linker
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cccgaattcg gg
                                                                     12
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<213> Rattus rattus
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Pro Asp Lys Gln Ala Ala Ala
<210> 11
<211> 33
<212> DNA
<213> Artificial sequence
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<223> Nucleic acid sequence from pBluescript SK-76.1 encoding rat GDNF
      N-terminus sequence
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                                                                     33
<210> 12
<211> 11
<212> PRT
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S-225N - 10 -

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<213> Rattus rattus
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Glu Arg Asn Arg Gln Ala Ala Ala Ala Ser Pro
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<212> DNA
<213> Artificial sequence
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       Oligonucleotide PCR primer DHD-26 to amplify DNA encoding rat GDN
       F polypeptide
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<213> Rattus rattus
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<221> misc_feature
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S-225N - 11 -

<220>

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Asp Lys Ile Leu Lys Asn Leu 1 5
<210> 15
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<213> Artificial Sequence
<220>
<223> Oligonucleotide primer PD1 to amplify rat GDNF probe
<400> 15
gacgggactc taagatg
                                                                     17
<210> 16
<211> 20
<212> DNA
<213> Artificial Sequence
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<223> Oligonucleotide primer DHD23 to amplify rat GDNF probe
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<223> N at position 3 is inosine
<220>
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<223> N at position 3 is inosine
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S-225N - 12 -

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ogagao		
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<220>		
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S-225N - 13 -

```
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<210> 21
<211> 46
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<400> 22
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cgcggtaccc agtctctgga gccgga
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<211> 33
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S-225N - 14 -

<220>

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<223> Synthetic adapter fragment for plasmid pCJ1
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gatctagaat tgtcatgttt gacagcttat cat
<210> 24
<211> 37
<212> DNA
<213> Artificial Sequence
<220>
      Polylinker sequence for plasmid pCJX1-1 with EcoRI and PSTI overh
<223>
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ttt Phe	aag Lys	atg Met	aag Lys	tta Leu -75	tgg Trp	gat Asp	gtc Val	gtg Val	gct Ala -70	gtc Val	tgc Cys	ctg Leu	gtg Val	ctg Leu -65	ctc Leu	100
cac His	acc Thr	gcg Ala	tcc Ser -60	gcc Ala	ttc Phe	ccg Pro	ctg Leu	ccc Pro -55	gcc Ala	ggt Gly	aag Lys	agg Arg	cct Pro -50	ccc Pro	gag Glu	148
gcg Ala	ccc Pro	gcc Ala -45	gaa Glu	gac Asp	cgc Arg	tcc Ser	ctc Leu -40	ggc Gly	cgc Arg	cgc Arg	cgc Arg	gcg Ala -35	ccc Pro	ttc Phe	gcg Ala	196
ctg Leu	agc Ser -30	agt Ser	gac Asp	tca Ser	aat Asn	atg Met -25	cca Pro	gag Glu	gat Asp	tat Tyr	cct Pro -20	gat Asp	cag Gln	ttc Phe	gat Asp	244
gat Asp -15	gtc Val	atg Met	gat Asp	ttt Phe	att Ile -10	caa Gln	gcc Ala	acc Thr	att Ile	aaa Lys -5	aga Arg	ctg Leu	aaa Lys	agg Arg -1	tca Ser 1	292
cca Pro	gat Asp	aaa Lys	caa Gln 5	atg Met	gca Ala	gtg Val	ctt Leu	cct Pro 10	aga Arg	aga Arg	gag Glu	cgg Arg	aat Asn 15	cgg Arg	cag Gln	340
gct Ala	gca Ala	gct Ala 20	gcc Ala	aac Asn	cca Pro	gag Glu	aat Asn 25	tcc Ser	aga Arg	gga Gly	aaa Lys	ggt Gly 30	cgg Arg	aga Arg	ggc Gly	388
cag Gln	agg Arg 35	ggc Gly	aaa Lys	aac Asn	cgg Arg	ggt Gly 40	tgt Cys	gtc Val	tta Leu	act Thr	gca Ala 45	ata Ile	cat His	tta Leu	aat Asn	436
gtc Val 50	act Thr	gac Asp	ttg Leu	ggt Gly	ctg Leu 55	ggc Gly	tat Tyr	gaa Glu	acc Thr	aag Lys 60	gag Glu	gaa Glu	ctg Leu	att Ile	ttt Phe 65	484
agg Arg	tac Tyr	tgc Cys	agc Ser	ggc Gly 70	tct Ser	tgc Cys	gat Asp	gca Ala	gct Ala 75	gag Glu	aca Thr	acg Thr	tac Tyr	gac Asp 80	aaa Lys	532
ata Ile	ttg Leu	aaa Lys	aac Asn 85	tta Leu	tcc Ser	aga Arg	aat Asn	aga Arg 90	agg Arg	ctg Leu	gtg Val	act Thr	gac Asp 95	aaa Lys	gta Val	580
gly	Gln	Ala	Cys	tgc Cys	Arg	Pro	Ile	gcc Ala	ttt Phe	Asp	Asp	Asp	Leu	tcg Ser	ttt Phe	628
tta Leu	gat Asp 115	Asp	aac Asn	ctg Leu	gtt Val	tac Tyr 120	cat His	att Ile	cta Leu	aga Arg	aag Lys 125	His	tcc Ser	gct Ala	aaa Lys	676
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<213> Homo sapiens

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Ala Val Cys Leu Val Leu Leu His Thr Ala Ser Ala Phe Pro Leu Pro
-70 -65 -60 -55

Ala Gly Lys Arg Pro Pro Glu Ala Pro Ala Glu Asp Arg Ser Leu Gly -50 -45 -40

Arg Arg Arg Ala Pro Phe Ala Leu Ser Ser Asp Ser Asn Met Pro Glu -35 -30 -25

Asp Tyr Pro Asp Gln Phe Asp Asp Val Met Asp Phe Ile Gln Ala Thr -20 -15 -10

Ile Lys Arg Leu Lys Arg Ser Pro Asp Lys Gln Met Ala Val Leu Pro
-5 -1 1 5 10

Arg Arg Glu Arg Asn Arg Gln Ala Ala Ala Ala Asn Pro Glu Asn Ser 15 20 25

Arg Gly Lys Gly Arg Gly Gln Arg Gly Lys Asn Arg Gly Cys Val 30 35 40

Leu Thr Ala Ile His Leu Asn Val Thr Asp Leu Gly Leu Gly Tyr Glu
45 50 55

Thr Lys Glu Glu Leu Ile Phe Arg Tyr Cys Ser Gly Ser Cys Asp Ala 60 65 70

Ala Glu Thr Thr Tyr Asp Lys Ile Leu Lys Asn Leu Ser Arg Asn Arg 75 80 85 90

Arg Leu Val Thr Asp Lys Val Gly Gln Ala Cys Cys Arg Pro Ile Ala 95 100 105

Phe Asp Asp Leu Ser Phe Leu Asp Asp Asn Leu Val Tyr His Ile 110 115 120

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<213> Rattus rattus

<220>

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Ala Glu Asp His Ser Leu Gly His Arg Arg Val Pro Phe Ala Leu Thr 35 40 45

Ser Asp Ser Asn Met Pro Glu Asp Tyr Pro Asp Gln Phe Asp Asp Val 50 . 55 60

Met Asp Phe Ile Gln Ala Thr Ile Lys Arg Leu Lys Arg Ser Pro Asp 65 70 75 80

Lys Gln Ala Ala Leu Pro Arg Arg Glu Arg Asn Arg Gln Ala Ala 85 90 95

Ala Ala Ser Pro Glu Asn Ser Arg Gly Lys Gly Arg Arg Gly Gln Arg 100 105 110

Gly Lys Asn Arg Gly Cys Val Leu Thr Ala Ile His Leu Asn Val Thr 115 120 125

Asp Leu Gly Leu Gly Tyr Glu Thr Lys Glu Glu Leu Ile Phe Arg Tyr 130 135 140

Cys Ser Gly Ser Cys Glu Ala Ala Glu Thr Met Tyr Asp Lys Ile Leu 145 150 155 160

Lys Asn Leu Ser Arg Ser Arg Arg Leu Thr Ser Asp Lys Val Gly Gln
165 170 175

S-225N - 18 -

Ala Cys Cys Arg Pro Val Ala Phe Asp Asp Asp Leu Ser Phe Leu Asp 180 185 190

Asp Ser Leu Val Tyr His Ile Leu Arg Lys His Ser Ala Lys Arg Cys 195 200 205

Gly Cys Ile 210

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<223> Human pre-pro GDNF

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Ala Glu Asp Arg Ser Leu Gly Arg Arg Arg Ala Pro Phe Ala Leu Ser 35 40 45

Ser Asp Ser Asn Met Pro Glu Asp Tyr Pro Asp Gln Phe Asp Asp Val 50 55 60

Met Asp Phe Ile Gln Ala Thr Ile Lys Arg Leu Lys Arg Ser Pro Asp 65 70 75 80

Lys Gln Met Ala Val Leu Pro Arg Arg Glu Arg Asn Arg Gln Ala Ala 85 90 95

Ala Ala Asn Pro Glu Asn Ser Arg Gly Lys Gly Arg Arg Gly Gln Arg 100 105 110

Gly Lys Asn Arg Gly Cys Val Leu Thr Ala Ile His Leu Asn Val Thr 115 120 125 S-225N - 19 -

Asp Leu Gly Leu Gly Tyr Glu Thr Lys Glu Glu Leu Ile Phe Arg Tyr 130 135 140

Cys Ser Gly Ser Cys Asp Ala Ala Glu Thr Thr Tyr Asp Lys Ile Leu 145 150 155 160

Lys Asn Leu Ser Arg Asn Arg Arg Leu Val Thr Asp Lys Val Gly Gln $$ 165 $$ 170 $$ 175

Ala Cys Cys Arg Pro Ile Ala Phe Asp Asp Leu Ser Phe Leu Asp 180 185 190

Asp Asn Leu Val Tyr His Ile Leu Arg Lys His Ser Ala Lys Arg Cys 195 200 205

Gly Cys Ile 210

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<213> Homo sapiens

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<223> Human pre-pro GDNF N-terminal fragment

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Ala Glu Asp Arg Ser Leu Gly Arg Arg Arg Ala Pro Phe Ala Leu Ser 35 40 45

Ser Asp 50